

WEST Search History

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DATE: Tuesday, April 06, 2004

Hide?	Set Name	Query	Hit Count
		<i>DB=PGPB,USPT,EPAB,JPAB,DWPI; PLUR=YES; OP=ADJ</i>	
<input type="checkbox"/>	L8	13 and 14	4
<input type="checkbox"/>	L7	16 and rhodamin\$6.ab.	6
<input type="checkbox"/>	L6	14 and therap\$6	139
<input type="checkbox"/>	L5	13 and 14L4	0
<input type="checkbox"/>	L4	12 near5 salt	485
<input type="checkbox"/>	L3	photodynamic and L2	329
<input type="checkbox"/>	L2	rhodamine\$	25309
<input type="checkbox"/>	L1	us-5556992-\$.did.	2

END OF SEARCH HISTORY

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(FILE 'HOME' ENTERED AT 14:53:13 ON 06 APR 2004)

FILE 'HCAPLUS' ENTERED AT 14:53:21 ON 06 APR 2004

L1 19 S GUIMOND M?/AU
L2 5 S MOLFINO N?/AU
L3 1784 S ROY D?/AU
L4 1805 S L1-3
L5 3 S L4 AND RHODAMINE
SELECT RN L5 2

FILE 'REGISTRY' ENTERED AT 14:54:53 ON 06 APR 2004

L6 6 S E1-6

FILE 'HCAPLUS' ENTERED AT 14:55:37 ON 06 APR 2004

L7 1 S L6 AND L5
L8 13213 S L6
L9 113 S L8 AND RHODAMINE
L10 1 S L9 AND PHOTODYNAMIC
L11 2 S L9 AND DIAGNOSIS
L12 5474 S L8(L)THU/RL
L13 26 S L9 AND L12
L14 14 S L13 AND PY<2000

FILE 'REGISTRY' ENTERED AT 16:01:17 ON 06 APR 2004

L15 5 S L6 NOT 59865-13-3

FILE 'HCAPLUS' ENTERED AT 16:04:54 ON 06 APR 2004

L16 1 S L15
L17 0 S L16 NOT L5

FILE 'REGISTRY' ENTERED AT 16:05:41 ON 06 APR 2004

L18 STR 333957-98-5
L19 0 S L18
L20 52360 S OC5-C6-C6/ES
L21 0 S L18 SSS SAM SUB=L20
L22 STR L18
L23 42 S L22
L24 794 S L22 FUL
SAVE L24 ROY072P/A
L25 0 S L18 SSS SAM SUB=L24
L26 STR L18
L27 0 S L26 SSS SAM SUB=L24
L28 12 S L26 SSS FUL SUB=L24
SAVE L28 ROY072S1/A

FILE 'HCAPLUS' ENTERED AT 16:11:51 ON 06 APR 2004

L29 24 S L28
L30 3946 S PHOTODYNAMIC THERAPY+PFT/CT
L31 5182 S "PHOTOSENSITIZERS (PHARMACEUTICAL)"+PFT/CT
L32 162992 S LYMPHOCYTE+PFT,NT/CT
L33 35145 S "TRANSPLANT AND TRANSPLANTATION"+PFT/CT
L34 59562 S IMMUNITY+PFT,NT/CT
L35 10647 S RHODAMIN?/OBI
L36 23 S L35 AND L30
L37 12 S L30(L)RHODAMIN?
L38 9 S L37 NOT L29
SELECT RN L38 4

FILE 'REGISTRY' ENTERED AT 16:23:29 ON 06 APR 2004

L39 10 S E7-16

FILE 'HCAPLUS' ENTERED AT 16:24:10 ON 06 APR 2004

L40 6 S L38 AND L39

L41 1 S L40 AND PATENT/DT
L42 45210 S L39

FILE 'REGISTRY' ENTERED AT 16:27:56 ON 06 APR 2004

L43 0 S 333957-95-2/CRN
L44 44 S L24 AND BR>1
L45 39 S L44 AND N=2
L46 32 S L45 NOT IN/ELS
L47 24 S L46 AND BR<4
L48 10 S L47 AND " 4,5-DIBROMO"
L49 0 S L44 AND L39
L50 44 S L44 AND L24
L51 75 S L24 AND BR/ELS
L52 0 S L51 AND L39
L53 5 S L39 AND BR/ELS
L54 0 S " BENZOIC ACID, 2-(3,6-DIETHYLAMINO-4,5-DIBROMO-9H-XANTHEN-9-
L55 307 S OC5-C6-C6/ES AND 46.150.18/RID AND N=2 AND BR/ELS
L56 65 S L55 AND BR=2
L57 46 S L56 AND NRS=2
L58 2 S L57 AND O=1
L59 31 S L57 AND O=3
L60 17 S L59 AND "4,5-DIBROMO"
L61 4 S " 2-(3,6-DIAMINO-4,5-DIBROMO-9H-XANTHEN-9-YL)"
L62 2 S "4,5-DIBROMO" AND "BIS(DIETHYLAMINO)"
L63 5 S L39 AND BR=2

FILE 'HCAPLUS' ENTERED AT 16:41:44 ON 06 APR 2004

L64 2 S L62
L65 2 S L63
L66 3 S L64-65
S L24 AND BR/ELS

FILE 'REGISTRY' ENTERED AT 17:06:41 ON 06 APR 2004

L67 1223588 S BR/ELS

FILE 'HCAPLUS' ENTERED AT 17:06:42 ON 06 APR 2004

FILE 'REGISTRY' ENTERED AT 17:06:49 ON 06 APR 2004

L68 75 S L24 AND BR/ELS
L69 32 S L68 AND "DIETHYL"
L70 26 S L69 AND N=2
L71 57 S L55 AND "BIS(DIETHYLAMINO)"
L72 57 S L71 AND N=2
L73 2 S L72 AND "4,5-DIBROMO"
L74 1 S L73 AND CL/ELS
L75 STR 333957-96-3
L76 0 S L75 FAM
L77 1 S L75 FAM FUL

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(FILE 'HOME' ENTERED AT 14:53:13 ON 06 APR 2004)

FILE 'HCAPLUS' ENTERED AT 14:53:21 ON 06 APR 2004

L1 19 S GUIMOND M?/AU
 L2 5 S MOLFINO N?/AU
 L3 1784 S ROY D?/AU
 L4 1805 S L1-3
 L5 3 S L4 AND RHODAMINE
 SELECT RN L5 2

FILE 'REGISTRY' ENTERED AT 14:54:53 ON 06 APR 2004

L6 6 S E1-6

FILE 'HCAPLUS' ENTERED AT 14:55:37 ON 06 APR 2004

L7 1 S L6 AND L5

=> d ibib abs hitstr ind

L7 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 2001:265273 HCAPLUS
 DOCUMENT NUMBER: 134:292146
 TITLE: Rhodamine derivatives for photodynamic
 diagnosis and treatment
 INVENTOR(S): Roy, Denis-Claude; Guimond, Martin
 ; Molfino, Nestor A.
 PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital
 Maisonneuve-Rosemont
 SOURCE: PCT Int. Appl., 60 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
BR 2000014135	A	20020521	BR 2000-14135	20001003
EP 1267931	A1	20030102	EP 2000-965683	20001003
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
JP 2003510372	T2	20030318	JP 2001-527823	20001003
PRIORITY APPLN. INFO.: US 1999-157790P P 19991005				
WO 2000-CA1142 W 20001003				

AB The present invention relates to the use of the photoactivable derivs. for the photodynamic treatment for the selective destruction and/or inactivation of immunol. reactive cells without affecting the normal cells and without causing systemic toxicity for the patient, wherein appropriate intracellular levels of said derivs. are achieved and irradiation of a suitable wavelength and intensity is applied. Examples are given of the selective phototoxicity of rhodamine derivs. against K562 cells, CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and human B cells. Immunol. disorders, including graft-vs-host disease are treated with photodynamic therapy.

IT 333957-97-4

RL: BAC (Biological activity or effector, except adverse); BPR (Biological

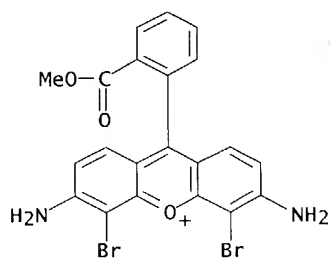
ROY 10/088,072

process); BSU (Biological study, unclassified); THU (Therapeutic use);
BIOL (Biological study); PROC (Process); USES (Uses)

(rhodamine derivs. for photodynamic diagnosis and treatment
of immunol. disorders)

RN 333957-97-4 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-,
bromide (9CI) (CA INDEX NAME)



● Br⁻

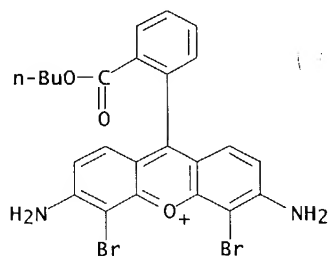
IT 333957-95-2 333957-96-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
(Uses)

(rhodamine derivs. for photodynamic diagnosis and treatment
of immunol. disorders)

RN 333957-95-2 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, bromide
(9CI) (CA INDEX NAME)

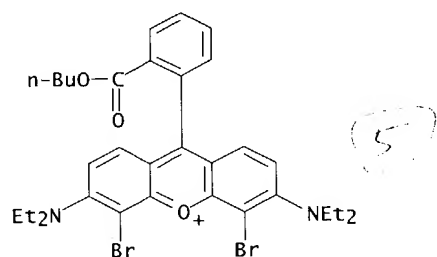


● Br⁻

RN 333957-96-3 HCAPLUS

CN Xanthylium, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-
, chloride (9CI) (CA INDEX NAME)

ROY 10/088,072



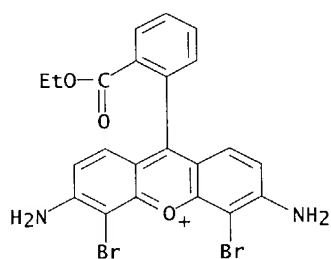
● Cl⁻

IT 333957-98-5 333957-99-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(rhodamine derivs. for photodynamic diagnosis and treatment
of immunol. disorders)

RN 333957-98-5 HCAPLUS

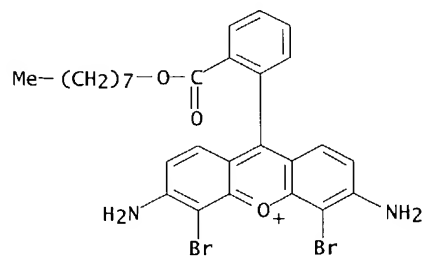
CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-, bromide
(9CI) (CA INDEX NAME)



● Br⁻

RN 333957-99-6 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-[(octyloxy)carbonyl]phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br⁻

IT 59865-13-3, Cyclosporin A

ROY 10/088,072

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(rhodamine derivs. for photodynamic diagnosis and treatment of immunol. disorders: effect of cyclosporin A on rhodamine cellular efflux)

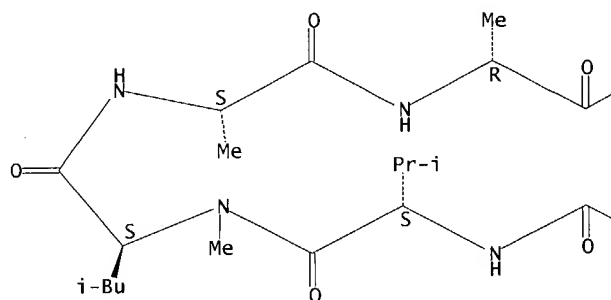
RN 59865-13-3 HCAPLUS

CN Cyclosporin A (9CI) (CA INDEX NAME)

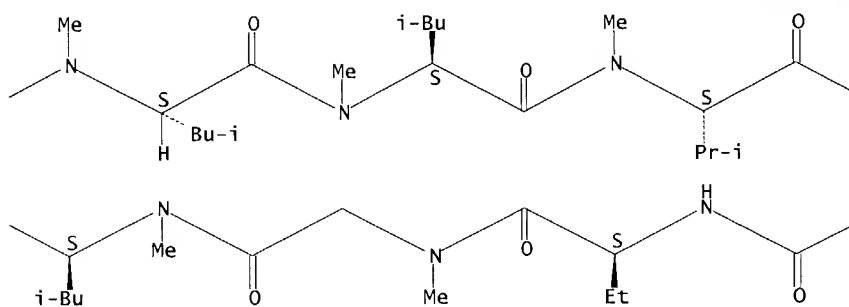
Absolute stereochemistry.

Double bond geometry as shown.

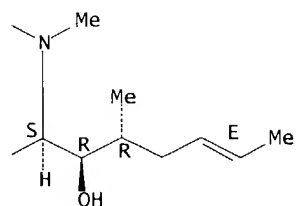
PAGE 1-A



PAGE 1-B



PAGE 1-C



IC ICM A61K041-00

CC 8-9 (Radiation Biochemistry)

Section cross-reference(s): 15

04/06/2004

Page 4

ST **rhodamine** deriv immunol disorder photodynamic therapy

IT Lymphocyte
(PHA-activated; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: phototoxicity against immunol. reactive cells)

IT Immunity
(disorder; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

IT Cytometry
(flow; photoactivatable **rhodamine** derivs. for evaluating transport mechanism of cells by flow cytometry)

IT Transplant and Transplantation
(graft-vs.-host reaction; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

IT Allergy
Autoimmune disease
Diagnosis
Photodynamic therapy
Photosensitizers (pharmaceutical)
Transplant rejection
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

IT B cell (lymphocyte)
CD4-positive T cell
CD8-positive T cell
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: phototoxicity against immunol. reactive cells)

IT Bone marrow
Hematopoietic precursor cell
Mononuclear cell (leukocyte)
Transplant and Transplantation
(**rhodamine** derivs. for photodynamic ex vivo treatment of hematopoietic cells)

IT 333957-97-4
RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

IT 333957-95-2 333957-96-3
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

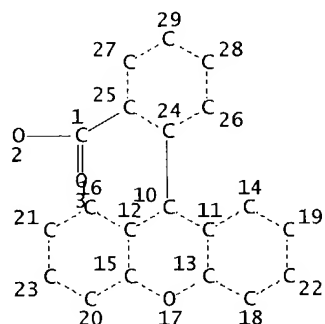
IT 333957-98-5 333957-99-6
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)

IT 59865-13-3, Cyclosporin A
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
(**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: effect of cyclosporin A on **rhodamine** cellular efflux)

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L22

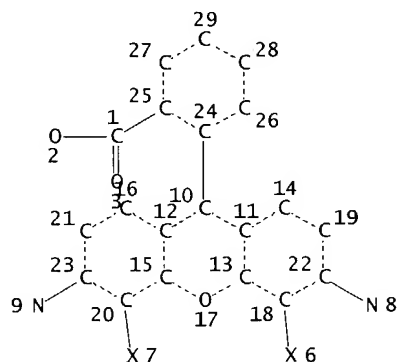
STR



NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RSPEC I
NUMBER OF NODES IS 23

STEREO ATTRIBUTES: NONE
L24 794 SEA FILE=REGISTRY SSS FUL L22
L26 STR



NODE ATTRIBUTES:
CONNECT IS X4 RC AT 8
CONNECT IS X4 RC AT 9
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RSPEC I
NUMBER OF NODES IS 27

STEREO ATTRIBUTES: NONE
L28 12 SEA FILE=REGISTRY SUB=L24 SSS FUL L26
L29 24 SEA FILE=HCAPLUS ABB=ON PLU=ON L28

=> d ibib abs hitstr 1-24

L29 ANSWER 1 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2003:749309 HCAPLUS
DOCUMENT NUMBER: 140:213073
TITLE: Ca2+-dependent and caspase-3-independent apoptosis

caused by damage in Golgi apparatus due to 2,4,5,7-tetrabromorhodamine 123 bromide-induced photodynamic effects

AUTHOR(S): Ogata, Maiko; Inanami, Osamu; Nakajima, Mihoko; Nakajima, Takayuki; Hiraoka, Wakako; Kuwabara, Mikiyori

CORPORATE SOURCE: Laboratory of Radiation Biology, Department of Environmental Veterinary Science, Graduate School of Veterinary Medicine, Hokkaido University, Sapporo, Japan

SOURCE: Photochemistry and Photobiology (2003), 78(3), 241-247
CODEN: PHCBAP; ISSN: 0031-8655

PUBLISHER: American Society for Photobiology

DOCUMENT TYPE: Journal

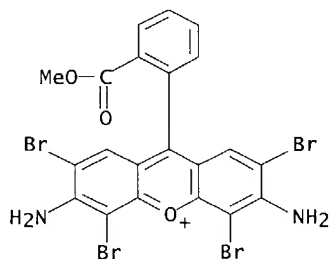
LANGUAGE: English

AB To clarify the role of the Golgi app. in photodynamic therapy-induced apoptosis, its signaling pathway was studied after photodynamic treatment of human cervix carcinoma cell line HeLa, in which a photosensitizer, 2,4,5,7-tetrabromorhodamine 123 bromide (TBR), was incorporated into the Golgi app. Laser scanning microscopic anal. of TBR-loaded HeLa cells confirmed that TBR was exclusively located in the Golgi app. HeLa cells incubated with TBR for 1 h were then exposed to visible light using an Xe lamp. Light of wavelength below 670 nm was eliminated with a filter. Morphol. observation of nuclei stained with Hoechst 33342 revealed that apoptosis of cells was induced by exposure to light. ESR spectrometry showed that light-exposed TBR produced both singlet oxygen (1O_2) and superoxide anion (O_2^-). Apoptosis induction by TBR was inhibited by pyrrolidine dithiocarbamate, an O_2^- scavenger, but not by NaN_3 , a quencher of 1O_2 . Furthermore, TBR-induced apoptosis was inhibited by aurointricarboxylic acid and $ZnCl_2$, which are known as inhibitors of DNase (DNase).gamma., and (acetoxymethyl)-1,2-bis(o-aminophenoxy)-ethane-N,N,N',N'-tetraacetic acid, a chelator of Ca^{2+} , but not by acetyl Asp-Glu-Val-Asp-aldehyde, an inhibitor of caspase-3. These results suggested that O_2^- was responsible for TBR-induced apoptosis, and Ca^{2+} -dependent and caspase-3-independent nuclease such as DNase .gamma. played an important role in apoptotic signaling triggered by Golgi dysfunction.

IT 623903-26-4, Tetrabromorhodamine 123
RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(tetrabromorhodamine 123; Golgi app., calcium, caspase, and DNase role in PDT-induced apoptosis in cervical carcinoma)

RN 623903-26-4 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br⁻

REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 2 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2003:226526 HCAPLUS

DOCUMENT NUMBER: 139:377312

TITLE: "Mitochondrial" photochemical drugs do not release toxic amounts of 102 within the mitochondrial matrix space

AUTHOR(S): Petrat, Frank; Pindiur, Stanislaw; Kirsch, Michael; de Groot, Herbert

CORPORATE SOURCE: Institut fur Physiologische Chemie, Universitaetsklinikum, Essen, D-45122, Germany

SOURCE: Archives of Biochemistry and Biophysics (2003), 412(2), 207-215

CODEN: ABBIA4; ISSN: 0003-9861

PUBLISHER: Elsevier Science

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Previously, we demonstrated that mitochondrial NAD(P)H is the primary target of singlet oxygen ($^{1}O_2$) generated by photoactivation of mitochondria-selective rhodamine derivs. Hence, local NAD(P)H oxidn./fluorescence decrease may be used to reveal the site of intracellular $^{1}O_2$ generation. Therefore, in addn. to the previously used tetramethylrhodamine methylester (TMRM), 2',4',5',7'-tetrabromorhodamine 123 bromide (TBRB) and rhodamine 123 (Rho 123), we tested here whether mitochondrial NAD(P)H of cultured hepatocytes is directly oxidized upon irradiation of different "mitochondrial" photosensitizers (Photofrin; protoporphyrin IX; Al(III) phthalocyanine chloride tetrasulfonic acid; meso-tetra(4-sulfonatophenyl)porphine dihydrochloride; Visudyne). In contrast to TMRM and Rho 123, which directly oxidized NAD(P)H upon irradiation, irradiation of intracellular TBRB and the photochem. drugs only indirectly affected mitochondrial NAD(P)H due to loss of mitochondrial integrity. In line with this result only TMRM and Rho 123 exclusively localized within the mitochondrial matrix. Due to these results it is doubtful whether real mitochondrial photosensitizers actually exist among the photochem. drugs applicable/used for photodynamic therapy.

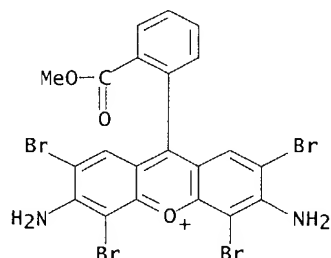
IT 623903-26-4

RL: PAC (Pharmacological activity); BIOL (Biological study)

(mitochondrial PDT photosensitizers do not release toxic amts. of $^{1}O_2$ within mitochondrial matrix space)

RN 623903-26-4 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)

● Br⁻

REFERENCE COUNT: 56 THERE ARE 56 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 3 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2002:777919 HCAPLUS

DOCUMENT NUMBER: 137:280622

TITLE: Halogenated rhodamine dye derivatives and their therapeutic applications
 INVENTOR(S): Habi, Abdelkrim; Gravel, Denis; Villeneuve, Luc; Forte, Jean-Pierre; Su, Hongsheng; Vaillancourt, Marc
 PATENT ASSIGNEE(S): Theratechnologies Inc., Can.
 SOURCE: PCT Int. Appl., 117 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002079183	A1	20021010	WO 2002-CA438	20020327
WO 2002079183	C1	20030220		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
EP 1276734	A1	20030122	EP 2002-708105	20020327
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BR 2002004489	A	20030401	BR 2002-4489	20020327
US 2003212126	A1	20031113	US 2003-297088	20030530
PRIORITY APPLN. INFO.:			CA 2001-2342675 A	20010402
			US 2001-822223 A	20010402
			WO 2002-CA438 W	20020327

OTHER SOURCE(S): MARPAT 137:280622

AB Bromo derivs. of rhodamine 110, rhodamine B, and rhodamine 6G and other halo rhodamine derivs. are useful as intermediates and as bactericides and antiviral agents and in the treatment of immunol. disorders. In an example, rhodamine B Me ester was dihydrogenated and then brominated and oxidized and treated with acetic acid to provide the purple acetate salt of 2,7-dibromorhodamine B Me ester.

IT 467232-05-9P

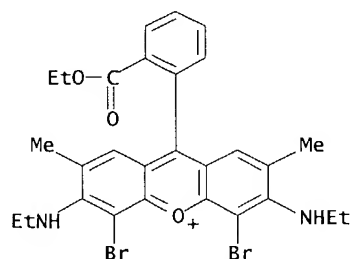
RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(green-red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)

RN 467232-05-9 HCAPLUS

CN Xanthylum, 4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-3,6-bis(ethylamino)-2,7-dimethyl-, bromide (9CI) (CA INDEX NAME)

ROY 10/088,072



● Br⁻

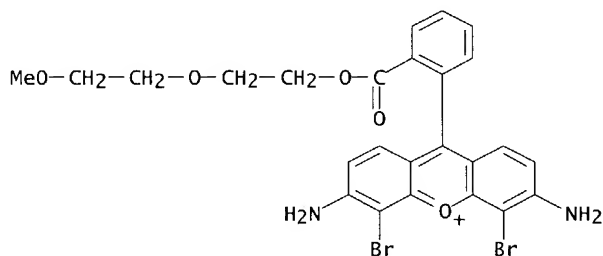
IT 467232-07-1P 467232-23-1P

RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)

RN 467232-07-1 HCAPLUS

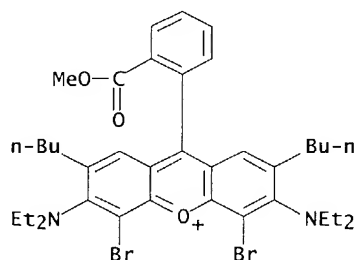
CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-[[2-(2-methoxyethoxy)ethoxy]carbonyl]phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br⁻

RN 467232-23-1 HCAPLUS

CN Xanthylum, 4,5-dibromo-2,7-dibutyl-3,6-bis(diethylamino)-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br⁻

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 4 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2002:541337 HCAPLUS

DOCUMENT NUMBER: 137:167817

TITLE: P-glycoprotein targeting: a unique strategy to selectively eliminate immunoreactive T cells

AUTHOR(S): Guimond, Martin; Balassy, Antonia; Barrette, Melanie; Brochu, Sylvie; Perreault, Claude; Roy, Denis Claude

CORPORATE SOURCE: Division of Hematology-Immunology, Maisonneuve-Rosemont Hospital Research Center, Department of Medicine, Universite de Montreal, Montreal, QC, Can.

SOURCE: Blood (2002), 100(2), 375-382

CODEN: BLOOAW; ISSN: 0006-4971

PUBLISHER: American Society of Hematology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB T lymphocytes have been found to harbor P-glycoprotein (Pgp) and to demonstrate modulation of its ion channel transporter function according to the state of activation of T lymphocytes. We hypothesized that cytotoxic chems. that are extruded by Pgp could be used to specifically eliminate immunoreactive T-cell populations. In this study, we evaluated the capacity of 4,5-dibromorhodamine Me ester (TH9402), a photosensitizer structurally similar to rhodamine, a dye transported by Pgp, and which becomes highly cytotoxic on activation with visible light to selectively deplete alloreactive T lymphocytes. Stimulation of T cells with mitogens or allogeneic major histocompatibility complex-mismatched cells resulted in the preferential retention of the TH9402 rhodamine-deriv. in activated T cells, both CD4+ and CD8+. Photodynamic cell therapy of TH9402-exposed T cells led to the selective elimination of immunoreactive T-cell populations. In addn., this treatment preserved resting T cells and their capacity to respond to third-party cells. Inhibition of Pgp enhanced cellular trapping of the dye in nonactivated T cells and resulted in their depletion after exposure to light. Targeting of Pgp-deficient cells may therefore represent an appealing strategy for the prevention and treatment of graft-vs.-host disease and other alloimmune or autoimmune disorders.

IT 174230-05-8, TH9402

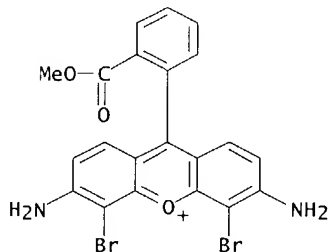
RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);

THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(P-glycoprotein targeting to selectively eliminate immunoreactive T cells)

RN 174230-05-8 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

REFERENCE COUNT: 78 THERE ARE 78 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 5 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2002:335895 HCAPLUS

DOCUMENT NUMBER: 137:75306

TITLE: Prevention of graft-versus-host disease while
preserving graft-versus-leukemia effect after
selective depletion of host-reactive T cells by
photodynamic cell purging processAUTHOR(S): Chen, Benny J.; Cui, Xiuyu; Liu, Congxiao; Chao,
Nelson J.CORPORATE SOURCE: Bone Marrow Transplantation Program, Duke University
Medical Center, Durham, NC, 27705, USASOURCE: Blood (2002), 99(9), 3083-3088
CODEN: BLOOAW; ISSN: 0006-4971

PUBLISHER: American Society of Hematology

DOCUMENT TYPE: Journal

LANGUAGE: English

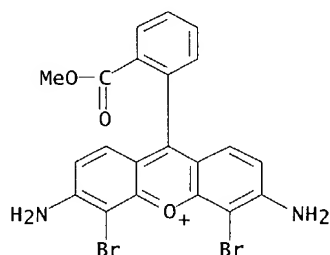
AB In this study, we investigated the possibility of selective depletion of donor alloantigen-specific T cells from C57BL/6 (H-2b) mice to prevent graft-vs.-host disease (GVHD). These cells were first activated with irradiated BALB/c (H-2d) host spleen cells in a 5-day mixed lymphocyte culture. Following this activation, a photoactive rhodamine deriv. called 4,5-dibromorhodamine 123 (TH9402), was added. This compd. is selectively retained in the mitochondria of activated host-reactive cells but not tumor-or third-party-specific resting cells. The treated cells were subsequently exposed to visible light (514 nm) to deplete the TH9402-enriched activated host-reactive cells. Treatment with photodynamic cell purging process (PDP) inhibited antihost responses measured by cytotoxic T lymphocytes (CTL) by 93%, and interferon- γ prodn. by 66%. By contrast, anti-BCL1 (BALB/c-origin leukemia/lymphoma) and anti-third-party C3H/HeJ (H-2k) responses were preserved. PDP-treated primed C57BL/6 cells were further tested in vivo. All lethally irradiated BALB/c mice inoculated with BCL1 cells and T-cell-depleted bone marrow cells developed leukemia by day +30, with 50% mortality by 100 days. All mice died of GVHD after addn. of 5 .times. 106 untreated primed C57BL/6 cells. However, addn. of same nos. of PDP-treated cells allowed 90% of the recipients to survive more than 100 days without detectable BCL1 tumor cells and free of GVHD. Moreover, PDP-treated primed C57BL/6 cells retained the ability to induce GVHD in the third-party C3H/HeJ mice. These data suggest that PDP can selectively deplete host alloantigen-specific T cells for GVHD prevention and immune and antileukemia function preserve.

IT 174230-05-8, TH9402

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(prevention of graft-vs.-host disease while preserving
graft-vs.-leukemia effect after selective depletion of host-reactive T
cells by photodynamic cell purging)

RN 174230-05-8 HCAPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-,
chloride (9CI) (CA INDEX NAME)

● Cl⁻

REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 6 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2001:265273 HCAPLUS

DOCUMENT NUMBER: 134:292146

TITLE: Rhodamine derivatives for photodynamic diagnosis and treatment

INVENTOR(S): Roy, Denis-Claude; Guimond, Martin; Molino, Nestor A.

PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital

Maisonneuve-Rosemont

SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
BR 2000014135	A	20020521	BR 2000-14135	20001003
EP 1267931	A1	20030102	EP 2000-965683	20001003
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
JP 2003510372	T2	20030318	JP 2001-527823	20001003
PRIORITY APPLN. INFO.:			US 1999-157790P P	19991005
			WO 2000-CA1142 W	20001003

AB The present invention relates to the use of the photoactivable derivs. for the photodynamic treatment for the selective destruction and/or inactivation of immunol. reactive cells without affecting the normal cells and without causing systemic toxicity for the patient, wherein appropriate intracellular levels of said derivs. are achieved and irradiation of a suitable wavelength and intensity is applied. Examples are given of the selective phototoxicity of rhodamine derivs. against K562 cells, CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and human B cells. Immunol. disorders, including graft-vs-host disease are treated with photodynamic therapy.

IT 333957-97-4

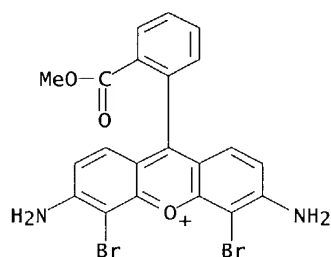
RL: BAC (Biological activity or effector, except adverse); BPR (Biological

ROY 10/088,072

process); BSU (Biological study, unclassified); THU (Therapeutic use);
BIOL (Biological study); PROC (Process); USES (Uses)
(rhodamine derivs. for photodynamic diagnosis and treatment of immunol.
disorders)

RN 333957-97-4 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-,
bromide (9CI) (CA INDEX NAME)



● Br⁻

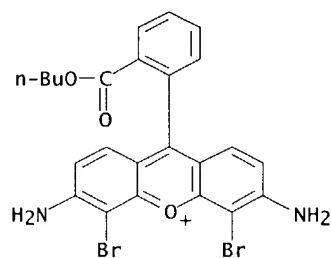
IT 333957-95-2 333957-96-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
(Uses)

(rhodamine derivs. for photodynamic diagnosis and treatment of immunol.
disorders)

RN 333957-95-2 HCAPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, bromide
(9CI) (CA INDEX NAME)

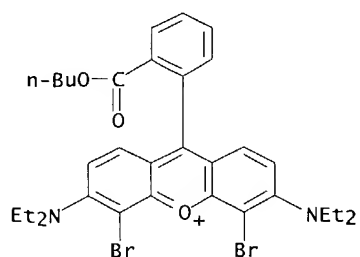


● Br⁻

RN 333957-96-3 HCAPLUS

CN Xanthylium, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-
, chloride (9CI) (CA INDEX NAME)

ROY 10/088,072



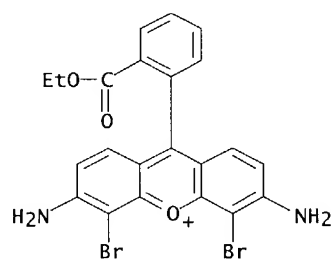
● Cl⁻

IT 333957-98-5 333957-99-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(rhodamine derivs. for photodynamic diagnosis and treatment of immunol.
disorders)

RN 333957-98-5 HCAPLUS

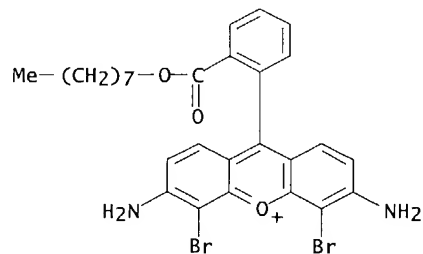
CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-, bromide
(9CI) (CA INDEX NAME)



● Br⁻

RN 333957-99-6 HCAPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-[(octyloxy)carbonyl]phenyl]-,
bromide (9CI) (CA INDEX NAME)



● Br⁻

REFERENCE COUNT:

8

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS

04/06/2004

Page 10

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 7 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2001:20398 HCAPLUS

DOCUMENT NUMBER: 134:204394

TITLE: Eradication of multiple myeloma and breast cancer cells by TH9402-mediated photodynamic therapy: implication for clinical ex vivo purging of autologous stem cell transplants

AUTHOR(S): Brasseur, N.; Menard, I.; Forget, A.; El Jastimi, R.; Hamel, R.; Molino, N. A.; Van Lier, J. E.

CORPORATE SOURCE: Department of Nuclear Medicine and Radiobiology, Faculty of Medicine, Université de Sherbrooke, Sherbrooke, QC, J1H 5N4, Can.

SOURCE: Photochemistry and Photobiology (2000), 72(6), 780-787
CODEN: PHCBAP; ISSN: 0031-8655

PUBLISHER: American Society for Photobiology

DOCUMENT TYPE: Journal

LANGUAGE: English

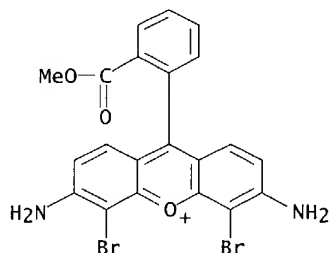
AB High-dose chemotherapy combined with autologous transplantation using bone marrow or peripheral blood-derived stem cells (PBSC) is now widely used in the treatment of hematol. malignancies as well as some solid tumors like breast cancer (BC). However, some controversial results were recently obtained in the latter case. The presence of malignant cells in the autograft has been assocd. with the recurrence of the disease, and purging procedures are needed to eliminate this risk. The aim of this study was to evaluate the potential of the photosensitizer 4,5-dibromorhodamine Me ester (TH9402), a dibrominated rhodamine deriv., to eradicate multiple myeloma (MM) and BC cell lines, while sparing more than 50% of normal pluripotential blood stem cells from healthy volunteers. The human BC MCF-7 and T-47D and MM RPMI 8226 and NCI-H929 cell lines were used to optimize the photodynamic purging process. Cell concn. and the cell suspension thickness as well as the dye and light doses were varied in order to eventually treat 1-2 L of apheresis. The light source consisted of two fluorescent scanning tubes emitting green light centered about 515 nm. The cellular uptake of TH9402 was measured during the incubation and washout periods and after photodynamic treatment (PDT) using spectrofluorometric anal. The limiting diln. assay showed that an eradication rate of more than 5 logs is obtained when using a 40 min incubation with 5-10 μ M dye followed by a 90 min washout period and a light dose of 5-10 J/cm² (2.8 mW/cm²) in all cell lines. Agitating the 2 cm thick cell suspension contg. 20 times. 10⁶ cells/mL during PDT was essential for maximal photoinactivation. Expts. on mobilized PBSC obtained from healthy volunteers showed that even more drastic purging conditions than those found optimal for maximal eradication of the malignant cell lines were compatible with a good recovery of hematopoietic progenitors cells. The absence of significant toxicity towards normal hematopoietic stem cells, combined with the 5 logs eradication of cancer cell lines induced by this procedure suggests that TH9402 offers an excellent potential as an ex vivo photodynamic purging agent for autologous transplantation in MM and BC treatment.

IT 174230-05-8, TH9402

RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
(myeloma and breast cancer cells eradication by TH9402-mediated photodynamic therapy: implication for clin. ex vivo purging of autologous stem cell transplants)

RN 174230-05-8 HCAPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

● Cl⁻

REFERENCE COUNT: 59 THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 8 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2000:911534 HCAPLUS

DOCUMENT NUMBER: 134:66121

TITLE: Compositions and methods for assaying subcellular conditions and processes using energy transfer for drug screening

INVENTOR(S): Dykens, James A.; Velicelebi, Gonul; Ghosh, Soumitra S.

PATENT ASSIGNEE(S): Mitokor, USA

SOURCE: PCT Int. Appl., 189 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

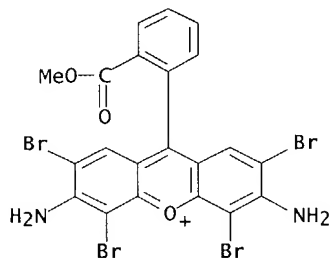
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000079274	A2	20001228	WO 2000-US17380	20000622
WO 2000079274	A3	20020110		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6323039	B1	20011127	US 1999-338122	19990622
US 6280981	B1	20010828	US 2000-514569	20000223
EP 1210596	A2	20020605	EP 2000-943119	20000622
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
JP 2003506014	T2	20030218	JP 2001-505191	20000622
PRIORITY APPLN. INFO.:				
			US 1999-140433P	P 19990622
			US 1999-338122	A 19990622
			US 2000-176383P	P 20000114
			WO 2000-US17380	W 20000622

AB The invention provides compns. and methods for monitoring subcellular compartments such as organelles by energy transfer techniques that do not require specific intermol. affinity binding events between energy transfer donor and energy transfer acceptor mols. pH. Provided are methods for assaying cellular membrane potential, including mitochondrial membrane potential, by energy transfer methodologies including fluorescence resonance energy transfer (FRET). Diagnostic and drug screening assays

are also provided.

IT 83796-96-7, Tetrabromo-rhodamine 123
 RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
 ANST (Analytical study); BIOL (Biological study); USES (Uses)
 (tetrabromorhodamine 123; compns. and methods for assaying subcellular
 conditions and processes using energy transfer for drug screening)
 RN 83796-96-7 HCAPLUS
 CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,
 chloride (9CI) (CA INDEX NAME)



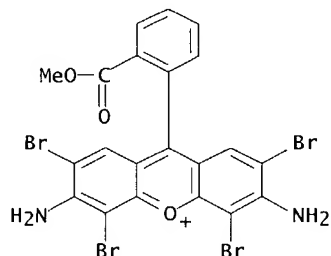
● Cl⁻

L29 ANSWER 9 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 2000:507144 HCAPLUS
 DOCUMENT NUMBER: 133:360498
 TITLE: Nonthermal ureteral tissue bonding: comparison of
 photochemical collagen crosslinking with thermal laser
 bonding
 AUTHOR(S): Merguerian, Paul A. M. D.; Pugach, Jeff L. M. D.;
 Lilge, Lothar D.
 CORPORATE SOURCE: Urology Div., Hospital for Sick Children, Univ. of
 Toronto, Toronto, ON, Can.
 SOURCE: Proceedings of SPIE-The International Society for
 Optical Engineering (1999), 3590(Lasers in Surgery:
 Advanced Characterization, Therapeutics, and Systems
 IX), 194-202
 CODEN: PSISDG; ISSN: 0277-786X
 PUBLISHER: SPIE-The International Society for Optical Engineering
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Because of difficulties with suture placement during minimally invasive
 procedures, many have sought alternative methods of creating tissue
 anastomoses. Although well studied, thermal laser tissue bonding has the
 potential of causing collateral thermal injury. Non-thermal tissue
 bonding agents, which cross-link proteins when activated with light, are
 currently being explored. We recently reported successful non-thermal
 bonding using tetrabromorhodamine (TBR). The bond was stronger than
 sutured repairs but weaker than laser thermal bonding. We currently
 report our ex-vivo experience with an alternate agent,
 riboflavin-5-phosphate and compare these results to thermal bonding and
 TBR. Successful ex vivo photochem. tissue welding with riboflavin of the
 rabbit ureter was achieved, without the generation of heat. Bond strength
 similar to that obtained with thermal welding was achieved.

IT 83796-96-7, Tetrabromo-rhodamine 123
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
 (Uses)
 (nonthermal ureteral tissue bonding: comparison of photochem. collagen
 crosslinking with thermal laser bonding)
 RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

REFERENCE COUNT: 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 10 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:570227 HCAPLUS

DOCUMENT NUMBER: 131:308414

TITLE: Ex vivo photodynamic purging in chronic myelogenous leukemia and other neoplasias with rhodamine derivatives

AUTHOR(S): Villeneuve, Luc

CORPORATE SOURCE: Theratechnologies Inc., Montreal, QC, H3B 1S6, Can.

SOURCE: Biotechnology and Applied Biochemistry (1999), 30(1), 1-17

CODEN: BABIEC; ISSN: 0885-4513

PUBLISHER: Portland Press Ltd.

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

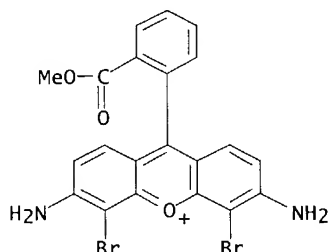
AB A review with 294 refs. Photodynamic therapy (PDT), a cancer treatment already used early in this century, has distinctive advantages over conventional chemotherapy, namely its often obsd. preferential accumulation in cancer cells and its low intrinsic toxicity. Aggressive therapeutic modalities using high doses of chemotherapy and/or radiation therapy are now commonplace treatments for leukemia, lymphoma and various non-haematol. malignancies. These intensive approaches have often been used in assocn. with hematopoietic-progenitor-cell support and have induced major responses and remissions in patients with relapsed and refractory diseases, ultimately contributing to improve the disease-free survival of patients with high risk. This has encouraged Theratechnologies, a Montreal-based pharmaceutical company, to develop photodynamic ex vivo purging procedures, including the development of new photosensitizers and irradiation devices for the safe eradication of neoplastic cells from autologous grafts. Our first specific objective, therefore, was to design, synthesize, purify and test photoactive rhodamine derivs. 4,5-Dibromorhodamine 123 (TH9402), a gas and phosphorus coating characteristic of an efficient scanning fluorescent source for extra-corporeal PDT using rhodamine derivs., was selected because of its photophys. properties, low toxicity and stability. TH9402 photodynamic-cell-therapy process conditions recognized as safe for normal human haematopoietic stem cells and progenitors demonstrated the efficacy of the purging procedure on various leukemias (including chronic-myelogenous-leukemia) as well as non-Hodgkin-leukemias and metastatic-breast-cancer cell lines.

IT 174230-05-8, TH 9402

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(TH 9402; ex vivo photodynamic purging in chronic myelogenous leukemia and other neoplasias with rhodamine derivs.)

RN 174230-05-8 HCAPLUS
 CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

REFERENCE COUNT: 292 THERE ARE 292 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

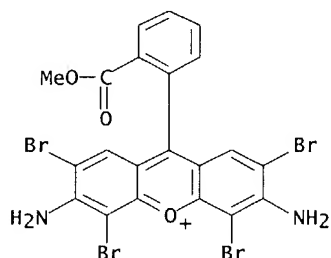
L29 ANSWER 11 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 1999:244672 HCAPLUS
 DOCUMENT NUMBER: 130:277634
 TITLE: Screening for oligonucleotide inhibitors of gene expression using conjugates with activatable reactive substances
 INVENTOR(S): Prescott, Catherine Denise
 PATENT ASSIGNEE(S): Smithkline Beecham Corporation, USA
 SOURCE: PCT Int. Appl., 21 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918116	A1	19990415	WO 1998-US21052	19981007
W: CA, JP, US				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 1023312	A1	20000802	EP 1998-952108	19981007
R: BE, CH, DE, DK, FR, GB, IT, LI, NL				
JP 2001519141	T2	20011023	JP 2000-514925	19981007
US 6387703	B1	20020514	US 2000-529095	20000406
PRIORITY APPLN. INFO.:		US 1997-61218P	P	19971007
		WO 1998-US21052	W	19981007

AB A method of screening for compds., particularly oligonucleotides, that modulate gene expression, particularly those which lower gene expression is described. The method uses a conjugate of the oligonucleotide and an activatable reactive group, such as a photosensitizing dye, preferably a compd. that generates reactive oxygen species. Target cells are incubated with the conjugate and the reactive group is activated and the effect on gene expression is assayed.

IT 83796-96-7D, Tetrabromo-rhodamine 123, derivs., conjugates with oligonucleotides
 RL: BUU (Biological use, unclassified); RCT (Reactant); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)
 (tetrabromorhodamine 123, photoactivatable inhibition of gene expression using; screening for oligonucleotide inhibitors of gene expression using conjugates with activatable reactive substances)

RN 83796-96-7 HCAPLUS
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 12 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:298888 HCAPLUS

DOCUMENT NUMBER: 129:51463

TITLE: [125I/127I/131I]Iodorhodamine: Synthesis, Cellular Localization, and Biodistribution in Athymic Mice Bearing Human Tumor Xenografts and Comparison with [99mTc]Hexakis(2-methoxyisobutylisonitrile)

AUTHOR(S): Harapanhalli, Ravi S.; Roy, Aloka M.; Adelstein, S. James; Kassis, Amin I.

CORPORATE SOURCE: Department of Radiology (Nuclear Medicine), Harvard Medical School, Boston, MA, 02115, USA

SOURCE: Journal of Medicinal Chemistry (1998), 41(12), 2111-2117

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

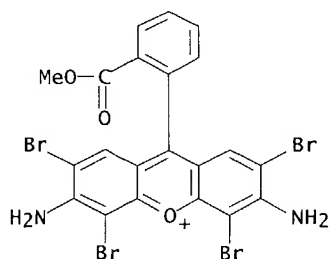
AB The synthesis of halogenated rhodamine (Rh) derivs. was carried out by controlling the stoichiometry of the halogenating agents, bromine and iodine monochloride. In the no-carrier-added synthesis of radioiodinated rhodamine 123, direct labeling of rhodamine 123 (Rh 123) with Na¹²⁵I/Na¹³¹I required the presence of the oxidant peracetic acid. ¹²⁵I/¹³¹I-Rh 123 was synthesized in modest yields (40-45%). HPLC purifn. sepd. Rh 123 from its mono- and diiodo derivs. Monohalogenation of Rh 123 did not alter the compd.'s ability to permeate viable cells and localize in mitochondria. ¹²⁵I/¹³¹I-Rh 123 was stable in serum in vitro but rapidly metabolized after i.v. injection into mice. Consequently, scintigraphy and biodistribution data reveal poor targeting of s.c. growing human tumor xenografts. The results are compared to those obtained following the administration of [99mTc]hexakis(2-methoxyisobutylisonitrile) which also did not image human tumor xenografts in nude mice.

IT 83796-96-7P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. and biodistribution of radioiodinated rhodamine 123 in tumor imaging)

RN 83796-96-7 HCAPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

● Cl⁻

REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 13 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1997:104125 HCAPLUS

DOCUMENT NUMBER: 126:183387

TITLE: Spectroscopic and photophysical investigations on the nature of localization of rhodamine-123 and its dibromo derivative in different cell lines

AUTHOR(S): Villeneuve, Luc; Pal, Prabir; Durocher, Gilles; Migneault, David; Girard, Denis; Giasson, Richard; Balassy, Antonia; Blanchard, Louise; Gaboury, Louis

CORPORATE SOURCE: Laboratoire de pathologie moléculaire, Département de pathologie, Université de Montréal, Montréal, QC, H3C 3J7, Can.

SOURCE: Journal of Fluorescence (1996), 6(4), 209-219

CODEN: JOFLEN; ISSN: 1053-0509

PUBLISHER: Plenum

DOCUMENT TYPE: Journal

LANGUAGE: English

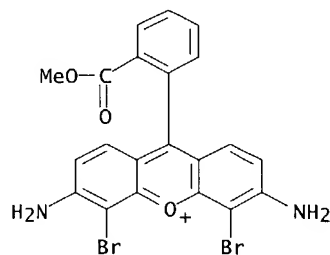
AB Steady-state and time-resolved spectroscopic properties of rhodamine-123 (rh123) and 4,5-dibromorhodamine Me ester (dbr123) bound to different cell lines are evaluated. Studies are also performed on the dye bound to extd. mitochondria. Results are compared with those obtained in homogeneous and microheterogeneous media. Results suggest that these dyes can specifically bind only with cell mitochondria. As a result of binding, excitation and emission spectra are red shifted by 10 to 12 nm. The fluorescence decay of these dyes bound to mitochondria shows two lifetimes. Values are about 4.0 and 2.0 ns for rh123 and about 1.9 and 0.5 ns for dbr123. Detailed global anal. of emission wavelength and dye concn. dependences of the fluorescence decay is performed. Results indicate that these dyes are bound to two different binding sites at mitochondria. The decay-assocd. fluorescence spectrum for the species corresponding to each binding site is recovered. Species 1, corresponding to the longer lifetime, is found to be more red shifted compared to species 2. The fluorescence of species 2 is heavily quenched. The origin of this quenching is explained in terms of resonance energy transfer between donor species 2 and acceptor species 1. The possible nature of the two binding sites is also discussed.

IT 174230-05-8

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (spectroscopic and photophys. investigations on the nature of localization of rhodamine-123 and its dibromo deriv. in different cell lines)

RN 174230-05-8 HCAPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

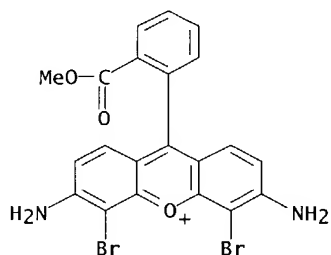
● Cl⁻

L29 ANSWER 14 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 1996:538169 HCAPLUS
 DOCUMENT NUMBER: 125:260967
 TITLE: Spectroscopic and photophysical properties of some new rhodamine derivatives in cationic, anionic and neutral micelles
 AUTHOR(S): Pal, P.; Zeng, H.; Durocher, G.; Girard, D.; Giasson, R.; Blanchard, L.; Gaboury, L.; Villeneuve, L.
 CORPORATE SOURCE: Laboratoire de photophysique moleculaire, Departement de chimie, Universite de Montreal, C.P. 6128, Succ. Centre-ville, Montreal, Que., H3C 3J7, Can.
 SOURCE: Journal of Photochemistry and Photobiology, A: Chemistry (1996), 98(1-2), 65-72
 CODEN: JPPCEJ; ISSN: 1010-6030
 PUBLISHER: Elsevier
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The spectroscopic and photophys. characterization of rhodamine 123 (dye 1), 4,5-dibromorhodamine Me ester (dye 2) and 4,5-dibromorhodamine Bu ester (dye 3) are reported in homogeneous media like water and some alcs. and also in microheterogeneous media; anionic sodium dodecylsulfate (SDS), cationic cetyltrimethylammonium bromide (CTAB) and neutral triton X-100 (TX) micelles. The selective biodistribution of these ionic drugs in tissues and membranes strongly influence their photosensitizing properties which have been part of our earlier studies. Results suggest that the hydrogen bonding capability of the amino end group lone pair of these dyes dominates in water. All these dyes interact with anionic SDS micelles. The interaction is mainly electrostatic in nature. At low SDS concns. (below c.m.c.), dye-SDS aggregate formation takes place. But above c.m.c. only the monomeric dye form is obsd. The penetration of dye 3 in SDS is a little less compared to dyes 1 and 2. Dyes 2 and 3 show a finite interaction with CTAB micelle unlike dye 1. With neutral TX micelles all the dyes form strong complexes. The fluorescence quantum yield (.PHI.F) of these three dyes in TX is lower. In time-resolved fluorescence expts., two lifetimes are obsd. The effects of the TX concn. on the fluorescence decay are measured. The decay assocd. spectra of dye 2 in TX are obtained by global compartmental anal. The dye-surfactant interaction mechanisms are also discussed.

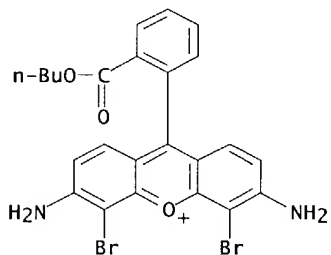
IT 174230-05-8 174230-06-9
 RL: PRP (Properties)
 (spectroscopic and photophys. properties of rhodamine derivs. in homogeneous media and micelles)

RN 174230-05-8 HCAPLUS
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

RN 174230-06-9 HCAPLUS
CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 15 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 1996:96151 HCAPLUS
DOCUMENT NUMBER: 124:197239
TITLE: Phototoxicity of some bromine-substituted rhodamine dyes: synthesis, photophysical properties and application as photosensitizers
AUTHOR(S): Pal, PRabir; Zeng, Hualing; Durocher, Gilles; Girard, Denis; Li, Tiechao; Gupta, Ajay K.; Giasson, Richard; Blanchard, Louise; Gaboury, Louis; et al.
CORPORATE SOURCE: Lab. Photophys. Mol., Univ. Montreal, Montreal, QC, Can.
SOURCE: Photochemistry and Photobiology (1996), 63(2), 161-8
CODEN: PHCBAP; ISSN: 0031-8655
PUBLISHER: American Society for Photobiology
DOCUMENT TYPE: Journal
LANGUAGE: English
AB The synthesis of some bromine-substituted rhodamine derivs., viz., 4,5-dibromorhodamine Me ester (dye 2) and 4,5-dibromorhodamine Bu ester (dye 3), are reported. These dyes were synthesized to promote a more efficient cancer cell photosensitizer for potential use in in vitro bone marrow purging in prepn. for autologous bone marrow transplantation. Spectroscopic and photophys. characterization of these dyes together with rhodamine 123 (dye 1) are reported in water, methanol, ethanol and also in a microheterogeneous system, sodium dodecyl sulfate. The possible mechanism of photosensitization os characterized in terms of singlet oxygen efficiency of these dyes. Singlet oxygen quantum yields for

bromine-substituted dyes are in the range of 0.3-0.5 depending on the solvent. For dye 1 no singlet oxygen prodn. is found. The photodynamic actions of these dyes in different cell lines are tested. It was found that dye 2 and dye 3 are efficient photosensitizers and mediate eradication of K562, EM2, myeloid cell lines (CML) and the SMF-AI rhabdomyosarcoma line.

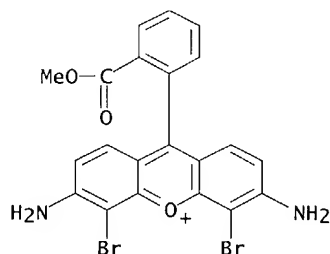
IT 174230-05-8P 174230-06-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(phototoxicity of some bromine-substituted rhodamine dyes: synthesis, photophys. properties and application in leukemia photosensitizations with laser radiation)

RN 174230-05-8 HCAPLUS

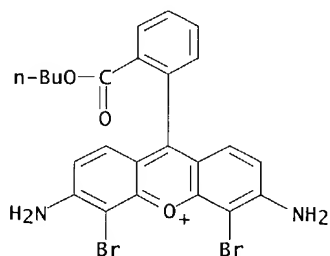
CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

RN 174230-06-9 HCAPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 16 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1995:709283 HCAPLUS

DOCUMENT NUMBER: 123:164113

TITLE: DMSO affects the efficiency of photolabeling of tetrabrominated rhodamine to collagen fibers.

AUTHOR(S): Jacques, Steven L.; Awazu, Kunio; Hasan, Tayyaba

CORPORATE SOURCE: M. D. Anderson Cancer Center, Univ. Texas, Houston, TX, 77030, USA

SOURCE: Proceedings of SPIE-The International Society for
Optical Engineering (1995), 2391(Laser-Tissue
Interaction VI), 232-7
CODEN: PSISDG; ISSN: 0277-786X

PUBLISHER: SPIE-The International Society for Optical Engineering

DOCUMENT TYPE: Journal

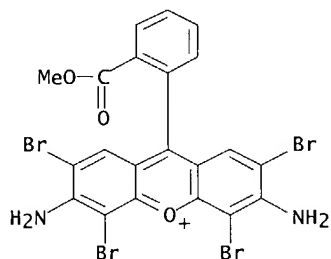
LANGUAGE: English

AB The ability to photolabel a compd., tetrabrominated rhodamine (TBR), onto
collagen gels was tested. The influence of DMSO on the efficiency of
photolabeling was detd. DMSO enhances the photolabeling presumably by
allowing TBR to become more closely assocd. to the collagen fibers such
that upon photon absorption which causes debromination to yield a radical,
the radical can covalently link to the collagen.

IT 83796-96-7
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
(Biological study); PROC (Process)
(DMSO affects the efficiency of photolabeling of tetrabrominated
rhodamine to collagen fibers)

RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,
chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 17 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1995:583719 HCAPLUS

DOCUMENT NUMBER: 123:78614

TITLE: Photodynamic therapy with cationic photosensitizers

AUTHOR(S): Kessel, David; Woodburn, Kathryn; Chang, Ck;
Henderson, Bw

CORPORATE SOURCE: Department Pharmacology, Wayne State University School
Medicine, Detroit, MI, USA

SOURCE: Proceedings of SPIE-The International Society for
Optical Engineering (1995), 2371, 334-8
CODEN: PSISDG; ISSN: 0277-786X

DOCUMENT TYPE: Journal

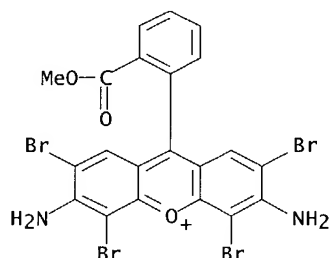
LANGUAGE: English

AB We characterized sites of photodamage catalyzed by two cationic
photosensitizers, tetrabromo-rhodamine 123 (TBR), which is recognized by
the multidrug transporter, and a monocationic porphyrin (MCP) which is
not. The transporter is an outward transport system assocd. with examples
of drug resistance. Irradn. of multidrug-resistant cells treated with TBR
resulted in highly-selective photodamage to the transporter site, while
MCP catalyzed nonspecific membrane damage to cells regardless of
transporter expression.

IT 83796-96-7, Tetrabromo-rhodamine 123
RL: BAC (Biological activity or effector, except adverse); BSU (Biological
study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
(Uses)
(photodynamic therapy with cationic photosensitizers)

ROY 10/088,072

RN 83796-96-7 HCAPLUS
CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 18 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1995:453091 HCAPLUS

DOCUMENT NUMBER: 122:285634

TITLE: Selective photodynamic inactivation of a multidrug transporter by a cationic photosensitizing agent

AUTHOR(S): Kessel, D; Woodburn, K

CORPORATE SOURCE: School of Medicine, Wayne State University, Detroit, MI, 48201, USA

SOURCE: British Journal of Cancer (1995), 71(2), 306-10

CODEN: BJCAAI; ISSN: 0007-0920

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We have characterized sites of photodamage catalyzed by the cationic photosensitizer tetrabromorhodamine 123, using P388 murine leukemia cells and a subline (P388/ADR) which has a multidrug resistance phenotype and hyperexpresses *mdr1* mRNA for P-glycoprotein. Fluorescence emission spectra were consistent with sensitizer localization in hydrophobic regions of the P388 cell, and in more aq. loci in P388/ADR. Subsequent irradiation resulted in photodamage to the P388 cells, resulting in loss of viability. In contrast, P388/ADR cells were unaffected except for an irreversible inhibition of P-glycoprotein, leading to enhanced accumulation of daunorubicin and rhodamine 123 and a corresponding increase in daunorubicin cytotoxicity. These results are consistent with the premise that substrates for P-glycoprotein are confined to membrane loci associated with the transporter, and indicate a very limited migration of cytotoxic photoproducts in a cellular environment.

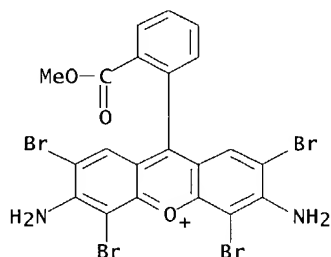
IT 83796-96-7, Tetrabromo-rhodamine 123

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(photodynamic inactivation of multidrug transporter in leukemia cells by cationic photosensitizer tetrabromorhodamine 123 with visible light)

RN 83796-96-7 HCAPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 19 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1993:18589 HCAPLUS

DOCUMENT NUMBER: 118:18589

TITLE: Mapping radiant energy distributions during laser irradiation of collagen phantoms by photolabeling with tetrabrominated rhodamine

AUTHOR(S): Jacques, Steven L.; Hasan, Tayyaba

CORPORATE SOURCE: Laser Biol. Res. Lab., Univ. Texas, Houston, TX, 77030, USA

SOURCE: Proceedings of SPIE-The International Society for Optical Engineering (1992), 1646(Proc. Laser-Tissue Interact. III, 1992), 219-26
CODEN: PSISDG; ISSN: 0277-786X

DOCUMENT TYPE: Journal

LANGUAGE: English

AB A method is proposed for mapping laser light distributions in gel phantoms by measurement of the distribution of a fluorescent compd. that has been photolabeled to the gel by the laser irradiance. A preliminary study of photolabeling by an argon laser using tetrabrominated rhodamine (TBR) was conducted in collagen gel phantoms to illustrate the feasibility of the method. A discussion of the basic quant. relationships for anal. of measurements is presented.

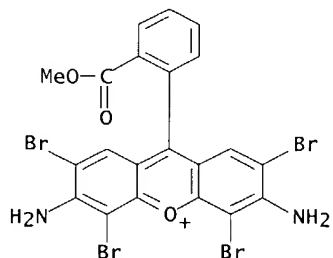
IT 83796-96-7

RL: BIOL (Biological study)

(photolabeling with, in laser radiation energy distribution mapping in collagen gel phantoms)

RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 20 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1992:146920 HCAPLUS

DOCUMENT NUMBER: 116:146920

TITLE: A test of the singlet oxygen mechanism of cationic dye photosensitization of mitochondrial damage

AUTHOR(S): Bunting, James R.

CORPORATE SOURCE: Baylor Res. Inst., Dallas, TX, 75226, USA

SOURCE: Photochemistry and Photobiology (1992), 55(1), 81-7

CODEN: PHCBAP; ISSN: 0031-8655

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Arom. cationic dyes have a potential as photo-chemotherapeutic agents because they are selectively concd. into the mitochondria of cancerous cells. The mechanism of cytophototoxicity has been proposed to be primarily due to dye sensitized photogeneration of highly toxic singlet oxygen (1O_2) at the mitochondria. This hypothesis was tested by measuring the relative phototoxicity of a collection of arom. cationic dyes towards respiring rat-liver mitochondria (RLM), upon addn. of 514 nm laser light. The effectiveness of dye photosensitization towards destruction of RLM function was assayed by its effect on the RLM membrane potential. Three phys. parameters of dye phototoxicity were independently measured and a relative phototoxicity calcd. assuming adherence of mechanism in the 1O_2 hypothesis. Quantum yields of dye-sensitized 1O_2 prodn. were estd., either from time-resolved luminescence measurements of photosensitized 1O_2 formed, or by comparing rates of photobleaching of 1O_2 trap; the relative partition of dye into mitochondrial lipid was detd. gravimetrically; and the optical d. of dye was detd. in a lipid like Triton X 100 micellar environment. Under the assumption of the 1O_2 hypothesis, these parameters were used to predict a relative phototoxicity which was compared with that obsd. For 12 of the 14 dyes investigated, the obsd. and predicted phototoxicities were linearly correlated ($r = 0.85$), suggesting support of the 10- and 1000-fold more potent than predicted, suggesting an addnl. factor at play in their phototoxicity.

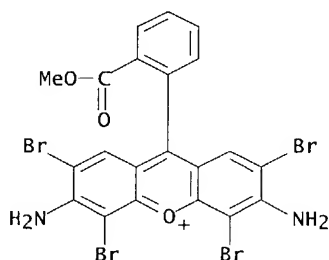
IT 83796-96-7, Tetrabromo-rhodamine 123

RL: BIOL (Biological study)

(photosensitization by, of liver mitochondria, singlet oxygen mechanism of evaluation in)

RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

● Cl⁻

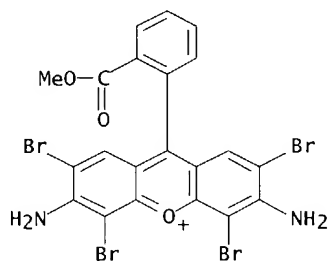
L29 ANSWER 21 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1989:473949 HCAPLUS

DOCUMENT NUMBER: 111:73949

TITLE: Rhodamine dyes as potential agents for photochemotherapy of cancer in human bladder carcinoma cells

AUTHOR(S): Shea, Christopher R.; Chen, Norah; Wimberly, Joanne; Hasan, Tayyaba
 CORPORATE SOURCE: Dep. Dermatol., Harvard Med. Sch., Boston, MA, 02114, USA
 SOURCE: Cancer Research (1989), 49(14), 3961-5
 CODEN: CNREA8; ISSN: 0008-5472
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The phototoxicity in vitro of rhodamine 123 and tetrabromorhodamine 123 (TBR) was compared in order to assess the photochemotherapeutic potential of these compds. Exposure to 514.5-nm radiation from an Ar ion laser caused phototoxicity in MGH-U1 bladder carcinoma cells previously treated with either dye at 10 .mu.M for 30 min. As assessed by colony formation and cellular morphol., TBR was markedly more phototoxic than rhodamine 123, reflecting increased intersystem crossing of TBR to the triplet manifold via spin-orbital coupling induced by the heavy Br atoms. Photoreactions of TBR very efficiently generated singlet O (102) in soln.; furthermore, irradiation of TBR-treated cells was significantly more toxic when performed in the presence of deuterium oxide, an enhancer of damage caused by 102. Retention of fluorescence in TBR-treated cells was enhanced by irradiation, indicating that a stable photoproduct may be formed in reaction with cellular components.
 IT 83796-96-7
 RL: PRP (Properties)
 (phototoxicity of, photochemotherapy of human bladder carcinoma in relation to)
 RN 83796-96-7 HCAPLUS
 CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 22 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 1989:208583 HCAPLUS
 DOCUMENT NUMBER: 110:208583
 TITLE: Phototoxicity of rhodamine dyes
 AUTHOR(S): Shea, Christopher R.; Chen, Norah; Hasan, Tayyaba
 CORPORATE SOURCE: Massachusetts Gen. Hosp., Harvard Med. Sch., Boston, MA, 02114, USA
 SOURCE: Proceedings of SPIE-The International Society for Optical Engineering (1989), 997(Adv. Photochemother.), 48-57
 CODEN: PSISDG; ISSN: 0277-786X
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Rhodamine-123 (R123) and tetrabromo-R123 (TBR) have been evaluated in vitro as potential photosensitizers with laser radiation. R123 localizes selectively in mitochondria of MGH-U1 bladder carcinoma cells exposed to 10 .mu.M R123 for 30 min, and under these conditions R123 is a weak photosensitizer. Incubation with R123 for longer times enhances its

phototoxicity, and is assocd. with a modification of its intracellular localization. TBR is .apprx.100-fold more phototoxic than R123, as assessed either by [3H]thymidine uptake or vital staining. Actively proliferating cells are more sensitive to either R123 or TBR phototoxicity than are plateau-phase, confluent cultures.

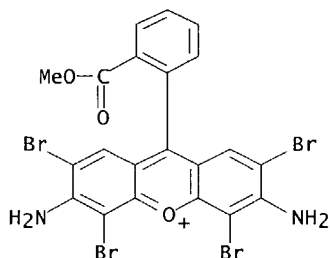
IT 83796-96-7

RL: PRP (Properties)

(phototoxicity of, to bladder carcinoma cells with laser radiation)

RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 23 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1983:211699 HCAPLUS

DOCUMENT NUMBER: 98:211699

TITLE: Triplet anisotropy decay measurements of DNA internal motion

AUTHOR(S): Hogan, Michael; Wang, Johnny; Austin, R. H.

CORPORATE SOURCE: Dep. Biochem. Sci., Princeton Univ., Princeton, NJ, 08540, USA

SOURCE: Ciba Foundation Symposium (1983), 93(Mobility Funct. Proteins Nucleic Acids), 226-45
CODEN: CIBSB4; ISSN: 0300-5208

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Triplet anisotropy decay techniques were used to measure the internal flexibility and overall rotational motions of DNA over a time range of 15 ns to 200 .mu.s. Nearly monodisperse DNA fragments with lengths varying from 65-600 base pairs were studied with the intercalating dye methylene blue as a triplet probe. The slow end-over-end tumbling of short DNA fragments (<165 base pairs) is as predicted for a rigid rod. A longer DNA fragment (600 base pairs) experiences slow segmental motions of its helix axis. At the earliest times, anisotropy decays more rapidly than expected for a rigid rod, suggesting that, when it is bound, methylene blue monitors fast internal motions of the helix. Since the rodlike end-over-end tumbling rules out fast bending motions (for short DNA fragments), the fast components of DNA anisotropy decay must be due to twisting motions of the helix, occurring with a time const. of .apprx.50 ns. The same techniques were used to measure the conformational flexibility of DNA in the nucleosome. It is concluded that, when the DNA helix is wrapped to form a nucleosome, it experiences substantial internal flexibility, occurring with a time const. of .apprx.30 ns. The amplitude and time-scale of this motion are similar to that seen in the uncomplexed DNA helix.

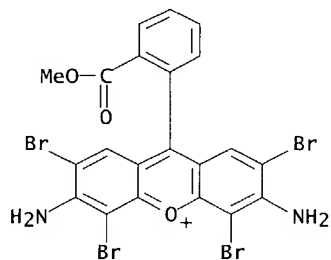
IT 83796-96-7

RL: BIOL (Biological study)

(triplet anisotropy decay of, in DNA, internal motions in relation to)

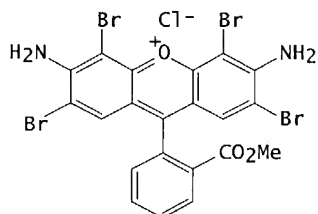
RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl⁻

L29 ANSWER 24 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN
 ACCESSION NUMBER: 1983:1816 HCAPLUS
 DOCUMENT NUMBER: 98:1816
 TITLE: DNA motions in the nucleosome core particle
 AUTHOR(S): Wang, J.; Hogan, M.; Austin, R. H.
 CORPORATE SOURCE: Dep. Biochem. Sci., Princeton Univ., Princeton, NJ, 08544, USA
 SOURCE: Proceedings of the National Academy of Sciences of the United States of America (1982), 79(19), 5896-900
 CODEN: PNASA6; ISSN: 0027-8424
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI



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AB Time-resolved triplet-state anisotropy decay techniques employing the intercalating agents, methylene blue and tetrabromorhodamine 123 chloride (I), the latter prep'd. by bromination of rhodamine 123, were used to measure the conformational flexibility of DNA in the chicken erythrocyte nucleosome. In a nucleosome, the DNA helix experiences substantial internal flexibility, which occurs with a time const. of .apprx.30 ns. The data can be fit well by a modified version of the Barkley-Zimm model for DNA motion, allowing only DNA twisting motions and the overall tumbling of the nucleosome. That fit yields a calcd. torsional rigidity equal to 1.8 .times. 10⁻¹⁹ erg-cm, a value equal to that measured for uncomplexed DNA. Such similarity suggests that large, fast twisting motions of the DNA helix persist, nearly unaltered, when DNA is wrapped to form a nucleosome.

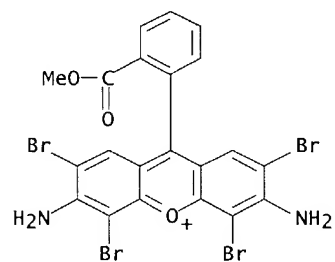
IT 83796-96-7P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (prepn. of, as DNA intercalating agent)

RN 83796-96-7 HCAPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,

ROY 10/088,072

chloride (9CI) (CA INDEX NAME)



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L39 10 SEA FILE=REGISTRY ABB=ON PLU=ON (13558-31-1/BI OR 177989-33-2
/BI OR 177989-34-3/BI OR 177989-35-4/BI OR 177989-36-5/BI OR
177989-37-6/BI OR 177989-38-7/BI OR 62669-70-9/BI OR 71-36-3/BI
OR 81-88-9/BI)
L62 2 SEA FILE=REGISTRY ABB=ON PLU=ON "4,5-DIBROMO" AND "BIS(DIETHY
LAMINO)"
L63 5 SEA FILE=REGISTRY ABB=ON PLU=ON L39 AND BR=2
L64 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L62
L65 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L63
L66 3 SEA FILE=HCAPLUS ABB=ON PLU=ON (L64 OR L65)

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L66 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2002:777919 HCAPLUS

DOCUMENT NUMBER: 137:280622

TITLE: Halogenated rhodamine dye derivatives and their
therapeutic applicationsINVENTOR(S): Habi, Abdelkrim; Gravel, Denis; Villeneuve, Luc;
Forte, Jean-Pierre; Su, Hongsheng; Vaillancourt, Marc

PATENT ASSIGNEE(S): Theratechnologies Inc., Can.

SOURCE: PCT Int. Appl., 117 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002079183	A1	20021010	WO 2002-CA438	20020327
WO 2002079183	C1	20030220		
W:		AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM		
RW:		GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG		
EP 1276734	A1	20030122	EP 2002-708105	20020327
R:		AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR		
BR 2002004489	A	20030401	BR 2002-4489	20020327
US 2003212126	A1	20031113	US 2003-297088	20030530
PRIORITY APPLN. INFO.:			CA 2001-2342675	A 20010402
			US 2001-822223	A 20010402
			WO 2002-CA438	W 20020327

OTHER SOURCE(S): MARPAT 137:280622

AB Bromo derivs. of rhodamine 110, rhodamine B, and rhodamine 6G and other halo rhodamine derivs. are useful as intermediates and as bactericides and antiviral agents and in the treatment of immunol. disorders. In an example, rhodamine B Me ester was dihydrogenated and then brominated and oxidized and treated with acetic acid to provide the purple acetate salt of 2,7-dibromorhodamine B Me ester.

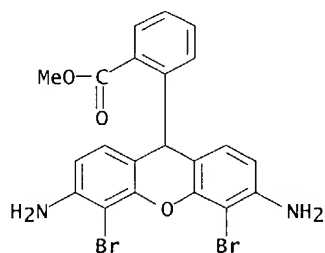
IT 177989-33-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(bacteriostatic agent; halogenated rhodamine dye derivs. and their therapeutic applications)

RN 177989-33-2 HCAPLUS

CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, methyl ester (9CI) (CA INDEX NAME)

ROY 10/088,072



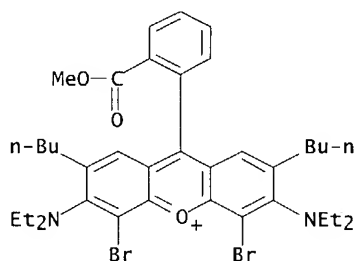
IT 467232-23-1P

RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)

RN 467232-23-1 HCAPLUS

CN Xanthylum, 4,5-dibromo-2,7-dibutyl-3,6-bis(diethylamino)-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



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REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L66 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2001:265273 HCAPLUS

DOCUMENT NUMBER: 134:292146

TITLE: Rhodamine derivatives for photodynamic diagnosis and treatment

INVENTOR(S): Roy, Denis-Claude; Guimond, Martin; Molfino, Nestor A.

PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital

Maisonneuve-Rosemont

SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,				

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ROY 10/088,072

SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

BR 2000014135 A 20020521 BR 2000-14135 20001003
EP 1267931 A1 20030102 EP 2000-965683 20001003

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL

JP 2003510372 T2 20030318 JP 2001-527823 20001003

PRIORITY APPLN. INFO.: US 1999-157790P P 19991005
WO 2000-CA1142 W 20001003

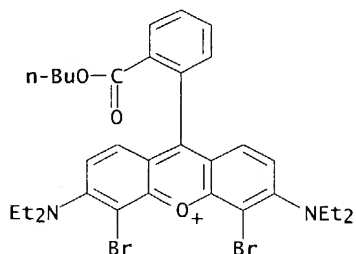
AB The present invention relates to the use of the photoactivable derivs. for the photodynamic treatment for the selective destruction and/or inactivation of immunol. reactive cells without affecting the normal cells and without causing systemic toxicity for the patient, wherein appropriate intracellular levels of said derivs. are achieved and irradiation of a suitable wavelength and intensity is applied. Examples are given of the selective phototoxicity of rhodamine derivs. against K562 cells, CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and human B cells. Immunol. disorders, including graft-vs-host disease are treated with photodynamic therapy.

IT 333957-96-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(rhodamine derivs. for photodynamic diagnosis and treatment of immunol. disorders)

RN 333957-96-3 HCAPLUS

CN Xanthylum, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-, chloride (9CI) (CA INDEX NAME)



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REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L66 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1996:379805 HCAPLUS

DOCUMENT NUMBER: 125:52522

TITLE: Novel rhodamine derivatives for photodynamic therapy of cancer and in vitro purging of the leukemias

INVENTOR(S): Gaboury, Louis; Giasson, Richard; Li, Tiechao; Gupta, Ajay Kumar; Villeneuve, Luc

PATENT ASSIGNEE(S): Universite De Montreal, Can.

SOURCE: PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

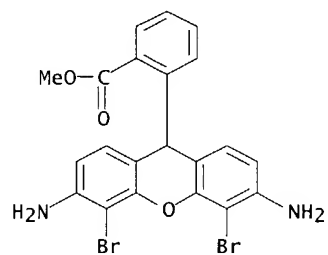
PATENT INFORMATION:

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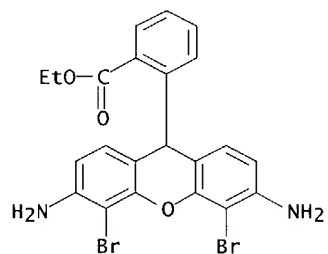
Page 3

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9607431	A1	19960314	WO 1995-CA485	19950816
W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TT				
RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
US 5556992	A	19960917	US 1994-300179	19940902
CA 2197435	AA	19960314	CA 1995-2197435	19950816
AU 9532488	A1	19960327	AU 1995-32488	19950816
AU 688100	B2	19980305		
EP 773794	A1	19970521	EP 1995-928907	19950816
EP 773794	B1	20010620		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
BR 9508779	A	19971223	BR 1995-8779	19950816
JP 10505349	T2	19980526	JP 1995-509057	19950816
AT 202286	E	20010715	AT 1995-928907	19950816
ES 2160173	T3	20011101	ES 1995-928907	19950816
PT 773794	T	20011228	PT 1995-95928907	19950816
US 5773460	A	19980630	US 1996-674247	19960701
GR 3036636	T3	20011231	GR 2001-401494	20010917
PRIORITY APPLN. INFO.:			US 1994-300179	A 19940902
			WO 1995-CA485	W 19950816
AB	The present invention relates to novel photoactivable rhodamine derivs. for enhancing high quantum-yield prodn. and singlet oxygen generation upon irradsn. with light while maintaining desirable differential retention of rhodamine between normal and cancer cells, said derivs. are selected from the group consisting of 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Me ester hydrochloride); 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Et ester hydrochloride); 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid octyl ester hydrochloride); 4,5-dibromorhodamine 110 Bu ester (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Bu ester hydrochloride); rhodamine B Bu ester (2-(6-Et amino-3-Et imino-3H-xanthen-9-yl)-benzoic acid Bu ester hydrochloride); and photoactivable derivs. thereof; whereby photoactivation of the derivs. induces cell killing while unactivated derivs. are substantially non-toxic to cells. Also, the present invention relates to the use of the photoactivable derivs. of the present invention for the photodynamic therapy of a cancer patient by destroying human cancer cells, wherein appropriate intracellular levels of the derivs. are achieved and irradsn. with light of a suitable wavelength is applied. The present invention also relates to a method for the photodynamic therapy of a patient suffering from leukemias, disseminated multiple myelomas or lymphomas.			
IT	177989-33-2 177989-34-3 177989-35-4 177989-36-5 177989-37-6 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (rhodamine derivs. for photodynamic therapy of cancer and leukemias)			
RN	177989-33-2 HCAPLUS			
CN	Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, methyl ester (9CI) (CA INDEX NAME)			

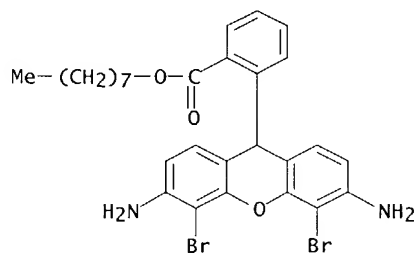
ROY 10/088,072



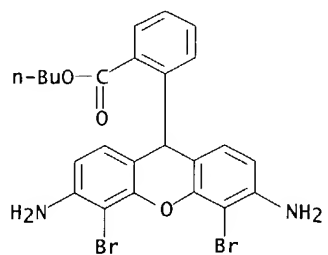
RN 177989-34-3 HCAPLUS
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, ethyl ester
(9CI) (CA INDEX NAME)



RN 177989-35-4 HCAPLUS
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, octyl ester
(9CI) (CA INDEX NAME)



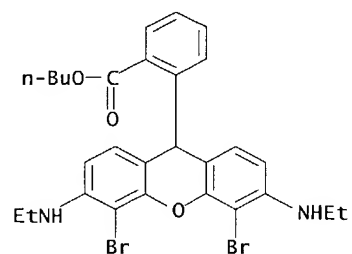
RN 177989-36-5 HCAPLUS
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, butyl ester
(9CI) (CA INDEX NAME)



ROY 10/088,072

RN 177989-37-6 HCAPLUS

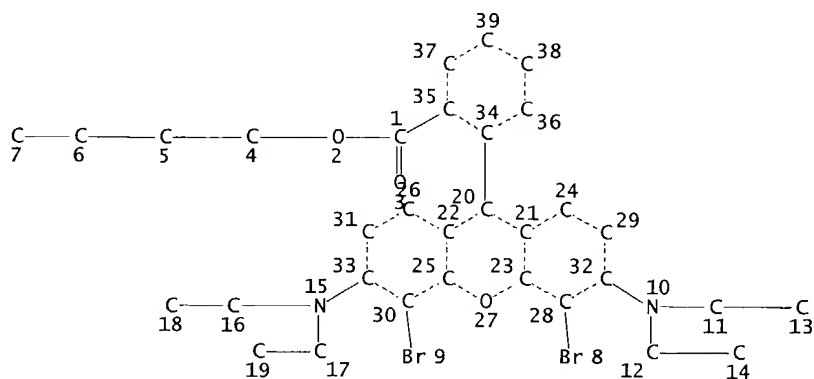
CN Benzoic acid, 2-[4,5-dibromo-3,6-bis(ethylamino)-9H-xanthen-9-yl]-, butyl ester (9CI) (CA INDEX NAME)



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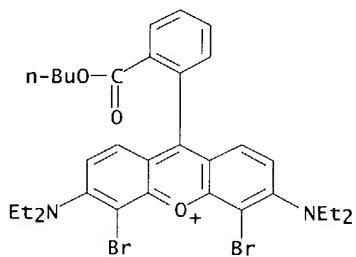
NODE ATTRIBUTES:
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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 39

STEREO ATTRIBUTES: NONE
L77 1 SEA FILE=REGISTRY FAM FUL L75

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L77 1 ANSWERS REGISTRY COPYRIGHT 2004 ACS on STN
IN Xanthylum, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-
, chloride (9CI)
MF C32 H37 Br2 N2 O3 . Cl



● Cl⁻

ALL ANSWERS HAVE BEEN SCANNED